

Claims Listing

1. (Original) An artificial LDL particle comprising an outer phospholipid monolayer and a solid lipid core, wherein the outer phospholipid monolayer comprises at least one apolipoprotein and the solid lipid core contains at least one therapeutic agent.
2. (Original) The artificial LDL particle of claim 1, wherein the at least one apolipoprotein is ApoE.
3. (Original) The artificial LDL particle of claim 2, wherein the at least one apolipoprotein is ApoE3.
4. (Original) The artificial LDL particle of claim 3, wherein the outer phospholipid monolayer further comprises one or more oxysterols and/or an additional apolipoprotein selected from the group consisting of ApoB and ApoE4.
5. (Original) The artificial LDL particle of claim 1, wherein the at least one therapeutic agent is selected from the group consisting of: amino acids, peptides, proteins, carbohydrates and lipids.
6. (Original) The artificial LDL particle of claim 1, wherein the at least one therapeutic agent is a conjugate formed between cholesterol and an agent selected from the group consisting of: amino acids, peptides, proteins, nucleic acids, carbohydrates and lipids.
7. (Original) The artificial LDL particle of claim 5, wherein the therapeutic agent is selected from the group consisting of: neurotrophic factors, growth factors, enzymes, antibodies, neurotransmitters, neuromodulators, antibiotics, antiviral agents, antifungal agents and chemotherapeutic agents.
8. (Original) The artificial LDL particle of claim 6, wherein the therapeutic agent is selected from the group consisting of: neurotrophic factors, growth factors, enzymes, neurotransmitters, neuromodulators, antibiotics, antiviral agents, antifungal agents and chemotherapeutic agents.

9. (Original) The artificial LDL particle of claim 1, wherein the outer phospholipid monolayer comprises phosphatidylcholine and at least one apolipoprotein.
10. (Original) The artificial LDL particle of claim 9, wherein the at least one apolipoprotein is ApoE.
11. (Original) The artificial LDL particle of claim 1, wherein the particle has a diameter between about 15 and 50 nm.
12. (Original) The artificial LDL particle of claim 1, wherein the particle has a diameter between about 20 and 30 nm.
13. (Original) The artificial LDL particle of claim 1, wherein the particle has a density between about 1.00 and 1.07 g/ml.
14. (Original) The artificial LDL particle of claim 1, wherein the particle has a density between about 1.02 and 1.06 g/ml.
15. (Original) The artificial LDL particle of claim 1, wherein the particle has a serum stability of at least two hours.
16. (Original) The artificial LDL particle of claim 1, wherein the particle is transported across the blood-brain barrier (BBB) by transcytosis.
17. (Original) The artificial LDL particle of claim 1, wherein the particle has at least a 3-fold greater uptake specificity for brain compared to liver.
18. (Original) The artificial LDL particle of claim 1, wherein the at least one therapeutic agent is a conjugate formed between cholesterol and adriamycin.
19. (Original) The artificial LDL particle of claim 1, wherein the at least one therapeutic agent is a conjugate formed between cholesterol and tetracycline.
20. (Original) The artificial LDL particle of claim 18, wherein the cholesterol and adriamycin of the conjugate are linked by an ester bond.

21. (Original) The artificial LDL particle of claim 19, wherein the cholesterol and tetracycline of the conjugate are linked by an ester bond.
22. (Original) An artificial LDL particle for delivery of an agent across the blood-brain barrier comprising an outer phosphatidylcholine monolayer, a solid lipid core comprising fatty acyl-cholesterol esters, and ApoE in the outer monolayer.
23. (Original) The artificial LDL particle of claim 22, wherein the solid lipid core further comprises cholesterol.
24. (Original) The artificial LDL particle of claim 22, wherein the ApoE in the outer monolayer is ApoE3.
25. (Original) A composition for delivery of an agent across the blood-brain barrier comprising the artificial LDL particle of claim 1 and a pharmaceutically acceptable carrier.
26. (Original) A composition for delivery of an agent across the blood-brain barrier comprising the artificial LDL particle of claim 4 and a pharmaceutically acceptable carrier.
27. (Original) A composition for delivery of an agent across the blood-brain barrier comprising the artificial LDL particle of claim 5 and a pharmaceutically acceptable carrier.
28. (Original) A conjugate comprising cholesterol linked to a therapeutic agent selected from the group consisting of: amino acids, peptides, proteins, nucleic acids, carbohydrates and lipids.
29. (Original) The conjugate of claim 28, wherein the therapeutic agent is selected from the group consisting of: neurotrophic factors, growth factors, enzymes, antibodies, neurotransmitters, neuromodulators, antibiotics, antiviral agents, antifungal agents and chemotherapeutic agents.

30. (Original) The conjugate of claim 29, wherein the therapeutic agent is adriamycin.
31. (Original) The conjugate of claim 30, wherein the adriamycin and cholesterol are linked by an ester linkage.
32. (Original) The conjugate of claim 29, wherein the therapeutic agent is tetracycline.
33. (Original) The conjugate of claim 32, wherein the tetracycline and cholesterol are linked by an ester linkage.
34. (Original) A method of producing an artificial LDL particle of claim 1 comprising the steps of: 1) suspending phospholipids containing conjugated or unconjugated therapeutic agent in a buffer solution; 2) sonicating the solution to form the outer phospholipid monolayer and solid lipid core; and 3) adding a solution comprising at least one apolipoprotein, wherein the apolipoprotein is incorporated into the outer phospholipid monolayer.
35. (Original) The method of claim 34, wherein the artificial LDL particles produced have a diameter between 10 and 50 nm.
36. (Original) A method for delivery a substance across the blood-brain barrier, said method comprising administering an effective amount of the composition of claim 25 to a mammal in need thereof.
37. (Original) A method for delivery a substance across the blood-brain barrier, said method comprising administering an effective amount of the composition of claim 26 to a mammal in need thereof.
38. (Original) A method for delivery a substance across the blood-brain barrier, said method comprising administering an effective amount of the composition of claim 27 to a mammal in need thereof.
39. (Original) A kit for delivering substances across the blood-brain barrier, said kit comprising a container containing the composition of claim 25 and instructions for use.

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40. (Original) A kit for delivering substances across the blood-brain barrier, said kit comprising a container containing the composition of claim 26 and instructions for use.
41. (Original) A kit for delivering substances across the blood-brain barrier, said kit comprising a container containing the composition of claim 27 and instructions for use.